

WHAT IS CLAIMED IS:

1. A tissue comprising linkers bonded to the tissue and a bridge molecule bonded between two or more of the linkers, wherein the linkers and the bridges are chemically different.
2. The tissue of claim 1 wherein the tissue comprises extracellular matrix selected from the group consisting of collagenous fibrils, GAG and elastin.
3. The tissue of claim 1 wherein the two linkers and the bridge bonded between the two linkers span a distance of between about 10 Angstroms and about 100 Angstroms.
4. The tissue of claim 1 wherein the two linkers and the bridge bonded between the two linkers span a distance of between about 15 Angstroms and about 50 Angstroms.
5. The tissue of claim 1 wherein the bridge is a single molecule.
6. The tissue of claim 1 wherein the bridge is reactive with modified tissue.
7. The tissue of claim 1 wherein the bridge comprises functional groups selected from the group

consisting of methylthio, thio, amine, alcohol, carboxyl and combinations thereof.

8. The tissue of claim 1 wherein the bridge comprises a hydrocarbon backbone.

9. The tissue of claim 1 wherein the linkers comprise monomers, dimers and oligomers.

10. The tissue of claim 1 wherein the linkers are active with respect to the tissue.

11. The tissue of claim 1 wherein the linkers comprise functional groups selected from the group consisting of aldehydes, epoxies, imide groups, photooxidative groups, enzymatically oxidative groups and combinations thereof.

12. The tissue of claim 1 wherein the linkers comprise crosslinking agents.

13. The tissue of claim 1 wherein the linker is selected from the group consisting of glutaraldehyde, triglycidyl amine and epoxy.

14. The tissue of claim 1 wherein a bioprosthetic device comprises the tissue.

15. The tissue of claim 14 wherein the bioprosthetic device is a heart valve prosthesis.
16. A method of crosslinking tissue comprising treating the tissue with a linker composition comprising linkers and a bridge composition comprising bridges wherein the linkers bond to the tissue and the bridges bond between two of the linkers, wherein the bridges and the linkers are chemically different.
17. The method of claim 16 wherein the tissue comprises proteins.
18. The method of claim 16 wherein the tissue is treated with the linker composition and the bridge composition simultaneously.
19. The method of claim 16 wherein the tissue is treated with the linker composition prior to addition of the bridge composition.
20. The method of claim 16 wherein the linker composition and the bridge composition are combined prior to treating the tissue.
21. The method of claim 16 wherein the linker composition comprises crosslinking agents.

22. The method of claim 16 wherein the concentration of the linkers in the linker composition is between about 0.0001 molar and about 1 molar.
23. The method of claim 16 wherein the concentration of the bridges in the bridge composition is between about  $1 \times 10^{-7}$  molar and about 1 molar.
24. The method of claim 16 wherein the tissue is treated with the linker composition and the bridge composition for between about 10 minutes and about one month.
25. The method of claim 16 wherein the tissue is treated with the linker composition and the bridge composition for between about 10 minutes and about 2 weeks.
26. The method of claim 16 wherein the bridges comprise multiple functional groups.
27. The method of claim 16 wherein the treatment of the tissue further comprises exposing the tissue to activators.
28. The method of claim 27 wherein the activators are selected from the group consisting of ultraviolet light, visible light and enzymes.

29. A method of bonding two molecules of linkers comprising adding bridge molecules, wherein the bridge molecules bond between the two of the linkers.

30. A composition comprising linkers and bridge molecules wherein the bridge molecules are bonded between two of the linkers, wherein the bridges and the linkers are chemically different.

31. The composition of claim 30 wherein the bridges comprise functional groups selected from the group consisting of methylthio, amine, alcohol, carboxyl and combinations thereof.

32. The composition of claim 30 wherein the linkers comprise functional groups selected from the group consisting of aldehydes, epoxies, imide groups, photooxidative groups, enzymatically oxidative groups and combinations thereof.

33. The composition of claim 30 wherein the concentration of the linkers in the composition is between about 0.0001 molar and about 1 molar and the concentration of the bridges is between about  $1 \times 10^{-7}$  molar and about 1 molar.

34. A tissue comprising bridge molecules, wherein the tissue is modified tissue and the bridge

molecules are bonded to two or more modified sites in the modified tissue.

35. The tissue of claim 34 wherein the modified sites comprise aldehyde groups.

36. A method of crosslinking tissue comprising treating modified tissue with a bridge composition comprising bridge molecules wherein the bridges bond to two or more modified sites in the modified tissue.

37. The method of claim 36 wherein the modified sites comprises aldehyde groups.